

# Intramedullary nail versus locking plate for treatment of proximal humeral fractures: A meta-analysis based on 1384 individuals

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### **Abstract**

**Objective:** The optimal surgical intervention on the treatment for proximal humeral fractures (PHFs) remains uncertain. The aim of this study was to evaluate clinical outcomes following fixation of PHFs by intramedullary nails or locking plates

**Methods:** The Cochrane Library, PubMed, EMBASE, China Knowledge Resource Integrated (CNKI), Chongqing VIP and Wanfang databases were systematically searched for studies published between January 01, 1996 and December 31, 2016 that investigated intramedullary nail vs. locking plate in the surgical treatment of PHFs. A meta-analysis examined incision length, blood loss, operation time, fracture healing time, Constant scores and post-operative complications. The methodological and evidence quality were also assessed by MINORS and GRADE system. **Results:** From the original 1024 references, 20 studies involving 1384 patients met the eligibility criteria. Analyses showed that intramedullary nails were superior to locking plates in incision length, peri-operative bleeding time, operation time and fracture healing time. However, there were no differences between treatments in Constant score or post-operative complications.

**Conclusion:** Although the evidence quality was poor, the results suggest that compared with locking plates, intramedullary nails may be a better choice for the repair of PHFs.

# **Keywords**

Intramedullary nail, locking plate, meta-analysis, proximal humeral fracture

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# Introduction

Proximal humeral fracture (PHF) is a common shoulder fracture of increasing prevalence which is common in elderly patients, many of whom have osteoporosis. 1-3 Recent estimates suggest that proximal humeral fractures account for 4-5% of all fractures and 20% of all fragility fractures.4,5 Although most PHFs do not require surgical intervention, surgical treatment is still necessary for comminuted or seriously displaced fractures including two, three, and four-part fractures. While, conventional plate and screw fixation has a high failure rate<sup>7</sup>, recent reports suggest that intramedullary nail and locking plates may provide good clinical outcome for unstable PHFs with rigid internal fixation.8

Although several studies have compared clinical outcomes following the use of intramedullary nail or locking plates for PHFs, the optimal management for the surgical treatment of these fractures remains controversial.9-28 A meta-analysis of eight studies conducted in 2015 that evaluated clinical outcomes and complications following insertion of locking plates or intramedullary nails for PHFs did not conclusively show their benefits and suggested more research is needed.<sup>29</sup> In addition, the study did not analyse the outcomes of the different types of PHFs according to their Neer classification.<sup>30</sup> Therefore, the purpose of the present meta-analysis was to evaluate evidence from studies that compared clinical outcomes following surgical treatment with intramedullary nail or locking plates in different types of PHFs.

### **Methods**

The Cochrane Library, PubMed, EMBASE, China Knowledge Resource Integrated (CNKI), Chongqing VIP and Wanfang databases were systematically searched for studies published between

January 01, 1996 and December 31, 2016 that investigated intramedullary nail vs. locking plate in the surgical treatment of PHFs. Key words/terms included: PHF; humeral neck fracture; PHN; PHILOS; intramedullary nail; proximal intramedullary nail; locking plate; locking proximal humerus plate. In addition, the reference lists of all included studies were checked for any potential additional publications. No restrictions were made on the publication language.

For a published report to be included in the meta-analysis, it had to fulfil the following criteria: (1) comparative randomized, controlled clinical trial or retrospective study of level III and above (Canadian Task Force on the Periodic Health Examination's Levels of Evidence),<sup>31</sup> that reported intramedullary nail vs. locking plate in the treatment of PHF; (2) reported at least one variable (e.g., incision length, blood loss, operation time, fracture healing time, Constant score<sup>32</sup> and/or postoperative complications); (3) follow-up time was at least 12 months. Single case reports, reviews, abstracts, letters, meeting proceedings and non-comparative were excluded.

Two reviewers [M.L., Y.H.W.] independently screened the papers from their titles and abstracts and selected relevant studies. The reviewers were not blinded to the journals, organizations, financial assistance, conflict of interest and/or researchers' information. The reviewers then assessed the full text to determine if the study should be included. Any discrepancy was resolved through discussion.

Three reviewers [M.L., M.Y., Y.P.Z.] independently extracted data from each study using a standard data extraction form. The following items were extracted: title; authors; date; country; study design; number of intramedullary nail and locking plate cases; number of female patients; Neer classification<sup>30</sup>; age, duration of

follow-up; clinical outcomes; level of evidence.<sup>31</sup>

The methodological quality of the randomized controlled was independently assessed by two authors [M.L., Y.H.W.] using the 'Cochrane collaboration's tool for assessing the risk of bias'. 33 Disagreements were resolved by discussion or consensus and a third author [M.Y] was the adjudicator when no consensus was reached. The tool covers seven domains: (1) random-sequence generation (selection bias); (2) allocation concealment (selection bias); (3) blinding of participants and personnel (performance bias); (4) blinding of outcome assessment (detection bias); (5) incomplete outcome data (attrition bias); (6) selective reporting (reporting bias); (7) other biases. All other studies were assessed by the same two authors using the methodological index for non-randomized studies (MINORS).<sup>34</sup> This is a validated instrument which is used to assess the quality of comparative or non-comparative controlled clinical trials. MINORS contain 12 items and each is scored as 0-2 (0= not reported; 1=reported but inadequate; 2= reported adequately). A MINORS score >12 was considered the level for inclusion in the meta-analysis.

The level of evidence quality was estimated according to the guidelines Grading of Recommendations, Assessment, Development, and Evaluation (GRADE).35 Although the GRADE working group acknowledges the importance of RCTs, it also recognizes circumstances in which observational studies generate high quality evidence of treatment effects. The GRADE approach applies a sequential assessment of the evidence quality and a subsequent judgment on the strength of the recommendations. The evidence grades were classified into four categories: (1) high grade (further research is unlikely to change confidence in the effect estimate); (2) moderate grade (further research is likely to

alter confidence in the effect estimate and may change the estimate); (3) low grade (further research is likely to alter confidence in the effect estimate significantly and to change the estimate); (4) very low grade (any effect estimate is uncertain). GRADEpro Guideline Development Tool (GDT) software version 3.6 for *Windows* was used to grade evidence and make recommendations.

# Statistical analyses

The meta-analysis was performed by two authors [M.L., Y.H.W.] using Review Manager (RevMan) [Computer program] Version 5.3. Copenhagen: The Nordic Cochrane Centre, The Cochrane Collaboration, 2014. A *P*-value <0.05 was considered to indicate statistical significance.

Prior to data analysis, heterogeneity was assessed according to Cochrane's  $\chi^2$  (Q) test and I<sup>2</sup> statistic. 36 No heterogeneity was defined as P > 0.1,  $I^2 = 0\%$ , low heterogeneity was P > 0.1,  $I^2 < 50\%$  and high heterogeneity was P < 0.1,  $I^2 > 50\%$ . For low statistical heterogeneity, a fixed effects model was applied to the data and for high heterogeneity a random effects model was used. Sensitivity analyses were performed by removing one study at a time and repeating the analyses. Weighted mean difference (WMD) and standardized mean difference (SMD) were used to measure the continuous variables and the relative risk (RR) was used for dichotomous  $(\pm 95\%)$ confidence intervals variables [Cis]). Publication bias was investigated with the use of a funnel plot graph. <sup>29</sup>

# Results

The literature search yielded an initial pool of 1024 references from which 20 articles ultimately met the eligibility criteria (Figure 1). 9-28 Twelve studies were

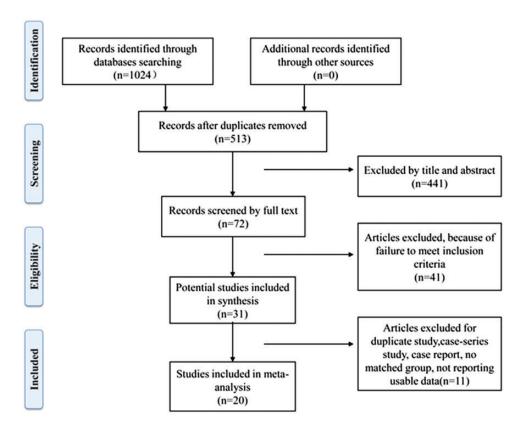


Figure 1. Flowchart of the study selection process.

performed in China, three in Germany, two in France, and one each in USA, Brazil and Spain (Table 1). Three studies were prospective and 17 were retrospective. Nine studies investigated Neer type II PHFs, one study investigated Neer type III and one type IV. Three other studies investigated Neer types II and III PHFs, two studies investigated Neer types III and IV PHFs, and four studies investigated Neer types II, III and IV PHFs.

In total, 1384 patients with PHFs were involved in the 20 studies, (631 patients in the intramedullary nail group and 753 in the locking plate group). Individual sample sizes ranged from 22 to 211 patients, the male/female ratio was similar across the studies and mean ages ranged from 49 to 76 years (Table 1).

Using the Cochrane collaboration's tool, <sup>33</sup> the methodological quality of the three randomised controlled trials <sup>26–28</sup> was assessed. The three studies had low risk of random sequence generation and allocation concealment (i.e., selection bias) but had high risk of blinding of outcomes (i.e., attrition and detection biases).

The mean MINOR score from the 17 non-randomised studies<sup>9–25</sup> was 16.7 (range, 15-17) which corresponded to a 70% score for the methodological quality of the included studies.

Five studies assessed incision length. Data from one study was separated into two parts according to Neer classification.<sup>25</sup> There was high heterogeneity between the results ( $I^2=97\%$ , P<0.05) and analysis showed that there was a statistically

Table I. Characteristics of the 20 studies included in the meta-analysis of intramedullary nail (IN) vs locking plate (LP) for the treatment of proximal humeral fractures.

Retrospective studies	Country	N (IN/LP)	% Female (IN/LP)	Classification (IN/LP)	age (y) (IN/LP)	(months) (IN/LP)	Study assessments
	China	26/28	89/59	II, 54	56/49	13/16	CS; ASES; peri-op bleeding time;
Gradl G et al, 2009 <sup>10</sup> Gern	Germany	9//9/	17/17	II, 52; III, 60: IV 40	63±16	12.8±0.4	Operations Complications
Matziolis D et al.2010 <sup>11</sup> Gern	Germany	11/11	64/64	II, 22	56±17/ 55+17	36	Complications
Ren S et al 2011 <sup>12</sup> China	na	21/15	67/73	II,6; III,13; IV,2/ II,4; III,8; IV,3	9∓09 9∓09	15	Incision length; peri-op bleeding time; operation time; healing
Lekic N et al, 2012 <sup>13</sup> USA	,	12/12	50/83	II, 24	69/09	12	time; Iveer score Complications; post-op
Konrad G et al, 2012 <sup>14</sup> Gern	Germany	58/153	81/74	III, 211	65±13/ 65+16	6 and 12	CS; Neer score; operation time;
Trepat A et al. 2012 <sup>15</sup>		15/14	47/57	11,29	65±12 65±21/ 68±17	6 and 12	Complications
Sheng S 2012 <sup>16</sup> China	na	22/24	54/54	II,46	67	15	Peri-op bleeding time; operation time; healing time; Neer score;
Boudard G et al. 2014 <sup>17</sup> France	ce	32/35	59/40	III,21; IV,9/ III.21: IV.12	64±16/ 50±18	12	CS; Complications
Xiaodong X et al, 2014 <sup>18</sup> China	na	14/24	64/71	II,8; III,5; IV,1/ II,7; III,1 IV,6	56±17/ 65±16	91	Incision length; operation time; healing time; CS; Complications; post-op
Wei C et al. 2014 <sup>19</sup> China	na	23/25	57/60	II, 48	55±6/ 58±6	22	Peri-op bleeding time; operation time; healing time; CS; Complications; post-op shaft angle

(continued)

Table 1. Continued.

Reference	Country	(IN/LP)	% Female (IN/LP)	Neer Classification (IN/LP)	Mean age (y) (IN/LP)	Follow-up (months) (IN/LP)	Study assessments
Yihan L et al. 2015 <sup>20</sup>	China	29/25	69/72	II,12; III,17/ II.11: III.14	74/76	3,6,12	CS; Complications
Hongbin B et al. 2015 <sup>21</sup>	China	26/34	54/56	11,60	67±5/ 66±7	16/15	Peri-op bleeding time; operation time; healing time; Neer score; ASES; post-op shaft angle; Complications
Gadea F et al. 2016 <sup>22</sup>	France	54/53	09/08	IV, 107	64/57	41/42	Incision length; CS; Complications
Haijun Y 2016 <sup>23</sup>	China	46/46	46/41	II,19; III,22; IV,5/ II,18; III,19; IV,9	63±11/ 62±11	12	Incision length; peri-op bleeding time; operation time; healing time; CS; Neer score; Complications
Ding Y et al. 2016 <sup>24</sup>	China	25/35	84/71	II,17; III,8/ II,20; III,15	74±3/ 74±3	12±3.0	Peri-op bleeding time; operation time; healing time; Neer score; Complications
Cheng G et al. 2016 <sup>25</sup>	China	54/54	52/56	III,36; IV,18/ III,32; IV,22	54±13/ 56±13	<u>4</u>	Incision length; peri-op bleeding time; operation time; healing time; CS; Complications; post-op shaft angle
Prospective studies Zhu Y et al. 2011 <sup>26</sup>	China	25/26	64/69	II, 51	55±17/ 51+20	12 and 36	CS; ASES; peri-op bleeding time;
Gracitelli M et al. 2016 <sup>27</sup>	Brazil	32/33	9//69	,16;    ,16/  ,16:    ,17	65±9/ 66+8	12	CS; Complications; post-op
Tian F. et al, 2016 <sup>28</sup>	China	30/30	37/37	11,60	26±5 56±5	na	Periop bleeding time; operation time; healing time; post-op shaft angle

Values are shown as *n*, mean or mean ± standard deviation IN, Intramedullary nail; LP, Locking plate; op, operative; CS, Constant score<sup>32</sup>; ASES, American Shoulder and Elbow Surgeons<sup>9</sup>; na, not available

significant difference in incision length between the intramedullary nail group and the locking plate group (Figure 2). The incision length for the intramedullary nail group was shorter compared with that for the locking plate group.

Six studies of Neer type II fractures and four of Neer type II, III and IV fractures (one study was separated into two parts according to Neer classification.  $^{25}$ ) reported peri-operative bleeding times. There was high heterogeneity between the studies ( $I^2=94\%$ , P<0.05) and analysis showed

that there was a statistically significant difference in bleeding times between the intramedullary nail group and the locking plate group (Figure 3). The blood loss for the intramedullary nail group was less than that for the locking plate group.

Six studies of Neer type II fractures and six of Neer type II, III and IV fractures (one study was separated into two parts according to Neer classification.  $^{25}$ ) reported operation time. There was high heterogeneity between the studies ( $I^2=78\%$ , P<0.05) and analysis showed that there was a

	Intrame	dullary	nail	locki	ng pl	ate		Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV. Random, 95% CI	IV. Random, 95% CI
Ren S et al. 2011	2.8	1.3	21	12.3	3.4	15	16.3%	-3.87 [-5.03, -2.72]	-
Xiaodong X et al. 2014	2.7	1.1	14	10.3	3.1	24	16.6%	-2.90 [-3.85, -1.95]	-
Gadea F et al. 2016	6.4	4.3	54	10.2	6	53	17.3%	-0.72 [-1.12, -0.33]	-
Haijun Y. 2016	3.1	0.4	46	11.7	2.5	46	16.8%	-4.76 [-5.58, -3.95]	-
Cheng G et al (III). 2016	3.1	1	36	12.5	2.3	32	16.5%	-5.35 [-6.39, -4.31]	-
Cheng G et al (IV). 2016	4.5	1.3	18	14	3.3	22	16.5%	-3.58 [-4.61, -2.54]	-
Total (95% CI)			189			192	100.0%	-3.51 [-5.30, -1.72]	•
Heterogeneity: Tau <sup>2</sup> = 4.77	; Chi <sup>2</sup> = 14	4.72, df	= 5 (P	< 0.0000	1);  2	= 97%			<del>' ' ' ' ' '</del>
Test for overall effect: Z = 3	3.85 (P = 0	.0001)	- 10		5.0				Intramedullary nail Locking plate

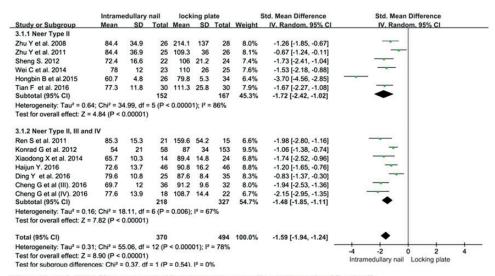
Cheng G et al. 2016 25 was separated into two parts according to Neer classification 30

**Figure 2.** Forest plot for meta-analysis of incision length between the intramedullary nail and locking plate groups using random effects model. Data from studies that assessed this parameter.

	Intram	edullary	nail	lock	king pla	ate		Std. Mean Difference	Std. Mean Difference			
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV. Random, 95% CI		IV. Random.	95% CI	
2.1.1 Neer Type II							_					
Zhu Y et al. 2008	214.1	137	26	382.1	200.6	28	9.6%	-0.96 [-1.52, -0.39]		-		
Zhu Y et al. 2011	214.1	140.2	25	382.1	200.6	26	9.6%	-0.95 [-1.53, -0.37]		-		
Sheng S. 2012	130	21.5	22	277	33.4	24	8.4%	-5.10 [-6.33, -3.86]		-		
Wei C et al. 2014	52	10	23	96	15	25	9.1%	-3.37 [-4.27, -2.46]				
Hongbin B et al.2015	180	13.5	26	300	10.6	34	6.8%	-9.92 [-11.83, -8.02]				
Tian F et al. 2016	51.6	9.8	30	96.5	15.2	30	9.2%	-3.47 [-4.28, -2.65]		_		
Subtotal (95% CI)			152			167	52.7%	-3.79 [-5.56, -2.02]		•		
Heterogeneity: Tau <sup>2</sup> = 4.61	; Chi2 = 1	36.05, d	f = 5 (P	< 0.000	01);  2 =	96%						
Test for overall effect: Z =	4.19 (P <	0.0001)										
2.1.2 Neer Type II, III and	IV											
Ren S et al. 2011	85.3	15.3	21	159.6	54.2	15	9.2%	-1.98 [-2.80, -1.16]		-		
Haijun Y. 2016	89.5	14.9	46	145.3	21.8	46	9.6%	-2.96 [-3.56, -2.36]		-		
Ding Y et al. 2016	82.4	27.5	25	149.4	26.5	35	9.4%	-2.46 [-3.14, -1.77]		-		
Cheng G et al (III). 2016	232.5	72.4	36	317.3	65.3	32	9.7%	-1.21 [-1.73, -0.69]		-		
Cheng G et al (IV). 2016	263.8	71.8	18	357.1	68.1	22	9.4%	-1.31 [-2.00, -0.62]		_		
Subtotal (95% CI)			146			150	47.3%	-1.98 [-2.70, -1.27]		•		
Heterogeneity: Tau <sup>2</sup> = 0.55	; Chi <sup>2</sup> = 2	4.10, df	= 4 (P <	0.0001	); I2 = 83	3%						
Test for overall effect: Z =	5.43 (P <	0.00001	)									
Total (95% CI)			298			317	100.0%	-2.85 [-3.73, -1.97]		•		
Heterogeneity: Tau <sup>2</sup> = 2.03	3; Chi <sup>2</sup> = 1	61.32, d	f = 10 (F	< 0.00	001); I <sup>2</sup>	= 94%			10	<del>-                                    </del>	<u> </u>	
Test for overall effect: Z =					,,				-10	-5 0 medullary nail Lo	5 cking plate	10

Cheng G et al. 2016 25 was separated into two parts according to Neer classification 30

**Figure 3.** Forest plot for meta-analysis of peri-operative bleeding loss for the intramedullary nail and locking plate groups using random effects model. Data from studies that assessed this parameter.



Cheng G et al. 2016 25 was separated into two parts according to Neer classification30

**Figure 4.** Forest plot for meta-analysis of operation time between intramedullary nail and locking plate groups using random effects model. Data from studies that assessed this parameter.

statistically significant difference in operation time between the intramedullary nail group and the locking plate group (Figure 4). The operation time for the intramedullary nail group was shorter than for the locking plate group.

Four studies of Neer type II fractures and five of Neer type II, III and IV fractures (one study was separated into two parts according to Neer classification. For properties according to the near type II, III and IV fractures according to Neer classification. For properties according to Neer classification ( $I^2 = 81\%$ ,  $I^2 = 81\%$ , and  $I^2 = 81\%$ ,

Three studies of Neer type II fractures and nine of Neer type II, III and IV fractures (one study was separated into two parts according to Neer classification.<sup>25</sup>) reported Constant scores. There was low heterogeneity between the studies

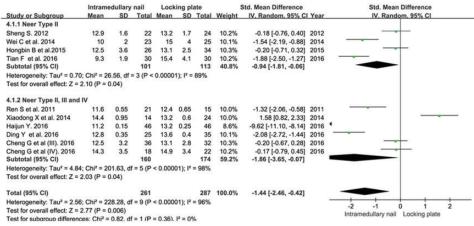
 $(I^2=30\%)$  and analysis showed that there was no statistically significant difference in Constant scores between treatment groups (Figure 6).

Seven studies of Neer type II fractures and eleven of Neer type II, III and IV fractures reported post-operative complications. There was low heterogeneity between the studies ( $I^2 = 40\%$ ) and analysis showed that there was no statistically significant difference in postoperative complications between treatment groups (Figure 7).

The above six outcomes were evaluated using the recommendations of the GRADE system<sup>35</sup> for level of evidence quality and the results were classed as 'very low grade' (Table 2).

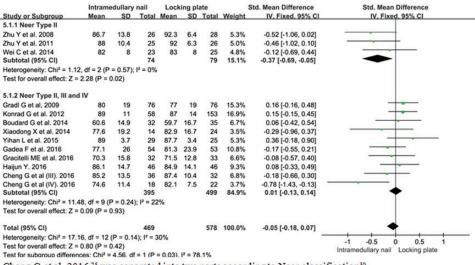
### Discussion

With the increasing aging population, PHFs have become one of the most common osteoporotic fractures seen in the elderly.<sup>4</sup> In recent years, intramedullary nails and locking plates have improved the



Cheng G et al. 2016 25 was separated into two parts according to Neer classification 30

**Figure 5.** Forest plot for meta-analysis of fracture healing time between intramedullary nail and locking plate groups using random effects model. Data from studies that assessed this parameter.

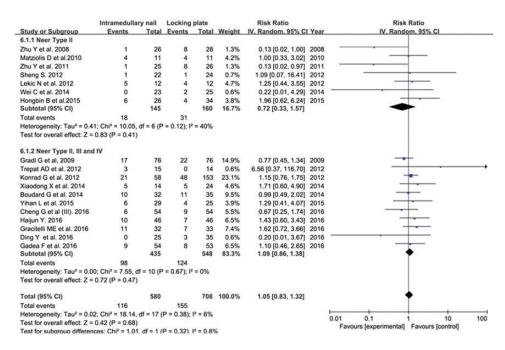


Cheng G et al. 2016 25 was separated into two parts according to Neer classification 30

**Figure 6.** Forest plot for meta-analysis of Constant score between the intramedullary nail and locking plate groups using a fixed effects model. Data from studies that assessed this parameter.

surgical outcome of PHFs and have gradually replaced rigid internal fixation by conventional plates. 9–28 However, there is currently no consensus in the literature on the best surgical technique for the different types of PHFs.

For routine Neer type II PHFs both intramedullary nails and locking plates have been shown to achieve good clinical results.<sup>27</sup> However, the use of these two methods for the treatment of complicated PHFs of type III and IV is a contentious



**Figure 7.** Forest plot for meta-analysis of post-operative complications between intramedullary nail and locking plate groups using a fixed effects model. Data from studies that assessed this parameter.

**Table 2.** Grading of Recommendations, Assessment, Development, and Evaluation (GRADE) evidence quality for each outcome.<sup>35</sup>

Quality assessment										
Outcomes	No. of studies	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Quality of evidence			
Incision length	5	serious	serious	none	none	none	Very low			
Peri-operative bleeding time	10	serious	serious	none	none	none	Very low			
Operation time	12	serious	serious	none	none	none	Very low			
Fracture healing time	9	serious	serious	none	none	none	Very low			
Constant score	12	none	serious	none	none	none	Very low			
Post-operative complications	18	none	serious	none	none	none	Very low			

issue. For example, one study reported intramedullary nail was not appropriate for Neer type III and IV PHFs because of the high rate of revision surgery required and low Constant score, 11 whereas, another study reported that intramedullary nail

insertion for complicated PHFs produced satisfactory results.<sup>37</sup> Because surgeons, particularly those in less developed countries, often select techniques according to their personal preferences, it is important to provide an evidence base so that they

can decide which is the best surgical method for the treatment of different types of PHF. To our knowledge, this current report is the first meta-analysis to use the GRADE system to evaluate results from studies comparing intramedullary nails and locking plates for PHFs. In addition, we evaluated data from different types of PHF which has not previously been considered.<sup>29</sup>

Results from this current meta-analysis of 20 published reports suggested that although the fixation of PHFs with intramedullary nails or locking plates produced similar outcomes in terms of Constant score and post-operative complications, the incision length, blood loss, operation time and fracture healing time were significantly shorter for the intramedullary nail group. Moreover, these results were independent of the type of PHF. One study suggested that the decreased operation time for the intramedullary nail procedure may be related to the minimal invasive approach of the nail implant.<sup>14</sup> However, stabilizing the PHF before implanting the nail required longer radiographic screening compared implanting the locking plate.<sup>14</sup> Another study suggested that the shorter operation time for implantation of the intramedullary nail resulted in less blood loss compared with implantation of the locking plate.<sup>38</sup>

Post-operative complications following repair of PHFs are of serious concern and may include, pain, infection, nonunion, osteonecrosis, secondary fracture, screw breakage, rotator cuff injury and impingement syndrome. While some studies have reported pain scores 2-3 times greater in the locking plate group than the intramedullary nail group, the studies have found more post-operative complications in the intramedullary nail group than in the locking plate group. Each procedure is associated with inherent difficulties. For example, although the insertion of a locking plate for PHFs can reduce the risk of

subacromial impingement syndrome and promote the recovery of shoulder function, due to its relatively larger volume, the procedure may cause iatrogenic injuries which may cause nonunion and avascular necrosis. In addition, although insertion of intramedullary nails can provide adequate fixation and favourable curative effects, defective surgery may damage important tendons and nerves and give rise to rotator cuff injuries and impingement syndromes. Moreover, intramedullary nails cannot be used in cases of severe osteoporosis. Some authors have suggested that the occurrence of post-operative complications following repair of PHFs are dependent on the experience and technique of the surgeon and not the surgical method. 15 Nevertheless, this current meta-analysis showed that, overall, there was no statistically significant difference between the surgical groups in the complication rate.

Using MINORS to assess the methodological quality of the included studies, a number of limitations of this meta-analysis were identified. For instance, poorly designed retrospective studies were more likely to suffer from various tapes of bias, especially publication bias including detection bias, selection bias, performance bias or attrition bias. In addition, due to the limited number of studies in the analysis, the potential difference between fixation with intramedullary nails and locking plates on Neer type III and IV PHFs could not be fully explored. Another limitation of the meta-analysis was that according to the GRADE recommendations for level of evidence quality, the results were classed as 'very low grade'.35

In summary, although the evidence quality was 'very low' in this meta-analysis because of clinical heterogeneity and limited information, the data tend to suggest that intramedullary nails may be superior to locking plated for the repair of PHFs. Further research is required and future

studies should include analysis of assessments at 3, 6, 12 and/or 24 months and follow-up after removal of the implants.

### **Author contributions**

M.L., and Y.H.W. conducted literature search and determined studies for exclusion and inclusion. M.L., M.Y., Y.P.Z., extracted data from the included studies, performed the meta-analysis, and drafted the manuscript. B.G.J., M.L., and P.X.Z. conceived the idea of the study, designed the study, and critically revised the manuscript for important intellectual content. All authors reviewed the paper and approved the final manuscript.

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# **Declaration of conflicting interests**

The authors declare that there are no conflict of interest.

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